0.16 gullens of 21/16/10/5 1. (CURRENTLY AMENDED) A compound of formula (I):

or stereoisomers or pharmaceutically acceptable salts thereof, wherein:

K is selected from  $CH_2$ ,  $CHR^5$  and  $CHR^6$ ;

L is selected from  $CH_2$ ,  $CHR^5$ ,  $CR^6R^6$  and  $CR^5R^6$ ;

J is selected from  $CH_2$ ,  $CHR^5$ ,  $CHR^{13}$ , and  $CR^5R^{13}$ ;

with the proviso:

at least one of K or L contains an R5;

Z is selected from O, S,  $NR^{1a}$ ,  $C(CN)_2$ ,  $CH(NO_2)$ , and CHCN;

 $R^{1a}$  is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl,  $CONR^{1b}R^{1b},\ OR^{1b},\ CN,\ NO_2,\ and\ (CH_2)_wphenyl;$ 

 $R^{1b}$  is independently selected from H,  $C_{1-3}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl;

E is  $-(C=0)-(CR^9R^{10})_v-(CR^{11}R^{12})-$ ,  $-(SO_2)-(CR^9R^{10})_v-(CR^{11}R^{12})-$ ,

Ring A is a  $C_{3-8}$  carbocyclic residue;

- $R^2$  is selected from H,  $C_{1-8}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, and a  $(CH_2)_r$ - $C_{3-10}$  carbocyclic residue substituted with 0-5  $R^a$ ;
- $R^b$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl;
- $R^c$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl;
- ${
  m R}^3$  is selected from  $({
  m CH}_2)_r {
  m N}({
  m CH}_3)_2$ , a  $({
  m CR}^3 {
  m 'R}^3 {
  m '})_r {
  m C}_{3-8}$  carbocyclic residue substituted with 0-5  ${
  m R}^{15}$ ; a  $({
  m CR}^3 {
  m 'R}^3 {
  m '})_r {
  m C}_{9-10}$  carbocyclic residue substituted with 0-4  ${
  m R}^{15}$ ; and a  $({
  m CR}^3 {
  m 'R}^3 {
  m '})_r 5-10$  membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  ${
  m R}^{15}$ ;

- $R^{3}$  and  $R^{3}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $(CH_2)_{r}C_{3-6}$  cycloalkyl, and phenyl;
- $R^5$  is selected from a  $(CR^5'R^5'')_t-C_{3-10}$  carbocyclic residue substituted with 0-5  $R^{16}$  and a  $(CR^5'R^5'')_t-5-10$  membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{16}$ ;
- $R^{5}$  and  $R^{5}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $(CH_2)_rC_{3-6}$  cycloalkyl, and phenyl;
- R6, at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8} \text{ alkenyl}, \ C_{2-8} \text{ alkynyl}, \ (CH_2)_r C_{3-6} \text{ cycloalkyl}, \\ (CF_2)_r CF_3, \ CN, \ (CH_2)_r NR^{6a}R^{6a'}, \ (CH_2)_r OH, \ (CH_2)_r OR^{6b}, \\ (CH_2)_r SH, \ (CH_2)_r SR^{6b}, \ (CH_2)_r C(0) OH, \ (CH_2)_r C(0) R^{6b}, \\ (CH_2)_r C(0) NR^{6a}R^{6a'}, \ (CH_2)_r NR^{6d}C(0)R^{6a}, \\ (CH_2)_r C(0) OR^{6b}, \ (CH_2)_r OC(0)R^{6b}, \ (CH_2)_r S(0)_p R^{6b}, \\ (CH_2)_r S(0)_2 NR^{6a}R^{6a'}, \ (CH_2)_r NR^{6d}S(0)_2 R^{6b}, \ and \\ (CH_2)_t phenyl \text{ substituted with 0-3 } R^{6c};$
- $R^{6a}$  and  $R^{6a'}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl substituted with 0-3  $R^{6c}$ ;
- $R^{6b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl substituted with 0-3  $R^{6c}$ ;

- $R^{6c}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6} \text{ cycloalkyl, Cl, F, Br, I, CN, NO}_2, (CF_2)_r CF_3, \\ (CH_2)_r OC_{1-5} \text{ alkyl, } (CH_2)_r OH, (CH_2)_r SC_{1-5} \text{ alkyl, and } \\ (CH_2)_r NR^{6d} R^{6d};$
- $R^{6d}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- with the proviso that when any of J, K, or L is CR<sup>6</sup>R<sup>6</sup> and R<sup>6</sup> is halogen, cyano, nitro, or bonded to the carbon to which it is attached through a heteroatom, the other R<sup>6</sup> is not halogen, cyano, or bonded to the carbon to which it is attached through a heteroatom;
- R<sup>9</sup>, is selected from H, C<sub>1-6</sub> alkyl, C<sub>2-8</sub> alkenyl, C<sub>2-8</sub> alkynyl, F, Cl, Br, I, NO<sub>2</sub>, CN, (CHR')<sub>r</sub>OH, (CH<sub>2</sub>)<sub>r</sub>OR<sup>9d</sup>, (CH<sub>2</sub>)<sub>r</sub>SR<sup>9d</sup>, (CH<sub>2</sub>)<sub>r</sub>NR<sup>9a</sup>R<sup>9a</sup>, (CH<sub>2</sub>)<sub>r</sub>C(O)OH, (CH<sub>2</sub>)<sub>r</sub>C(O)R<sup>9b</sup>, (CH<sub>2</sub>)<sub>r</sub>C(O)NR<sup>9a</sup>R<sup>9a</sup>, (CH<sub>2</sub>)<sub>r</sub>NR<sup>9a</sup>C(O)R, (CH<sub>2</sub>)<sub>r</sub>NR<sup>9a</sup>C(O)H, (CH<sub>2</sub>)<sub>r</sub>C(O)OR<sup>9b</sup>, (CH<sub>2</sub>)<sub>r</sub>OC(O)R<sup>9b</sup>, (CH<sub>2</sub>)<sub>r</sub>OC(O)NR<sup>9a</sup>R<sup>9a</sup>, (CH<sub>2</sub>)<sub>r</sub>NR<sup>9a</sup>C(O)OR<sup>9b</sup>, (CH<sub>2</sub>)<sub>r</sub>S(O)<sub>p</sub>R<sup>9b</sup>, (CH<sub>2</sub>)<sub>r</sub>S(O)<sub>2</sub>NR<sup>9a</sup>R<sup>9a</sup>, (CH<sub>2</sub>)<sub>r</sub>NR<sup>9a</sup>S(O)<sub>2</sub>R<sup>9b</sup>, C<sub>1-6</sub> haloalkyl, a (CH<sub>2</sub>)<sub>r</sub>-C<sub>3-10</sub> carbocyclic residue substituted with 0-5 R<sup>9c</sup>, and a (CH<sub>2</sub>)<sub>r</sub>-5-10 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3 R<sup>9c</sup>;

- $R^{9a}$  and  $R^{9a'}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r$ - $C_{3-10}$  carbocyclic residue substituted with 0-5  $R^{9e}$ , and a  $(CH_2)_r$ -5-10 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{9e}$ ;
- alternatively, R<sup>9a</sup> and R<sup>9a</sup>, along with the N to which they are attached, join to form a 5-6 membered heterocyclic system containing 1-2 heteroatoms selected from NR<sup>9g</sup>, O, and S and optionally fused with a benzene ring or a 6-membered aromatic heterocycle;
- $R^{9b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r$ - $C_{3-6}$  carbocyclic residue substituted with 0-2  $R^{9e}$ , and a  $(CH_2)_r$ -5-6 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{9e}$ ;

- $(CH_2)_rNR^{9f}S(O)_2R^{9b}$ , and  $(CH_2)_r$ phenyl substituted with 0-3  $R^{9e}$ ;
- $R^{9d}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  alkenyl,  $C_{3-6}$  alkynyl, a  $C_{3-10}$  carbocyclic residue substituted with 0-3  $R^{9c}$ , and a 5-6 membered heterocyclic system containing 1-4 heteroatoms selected from the group consisting of N, O, and S substituted with 0-3  $R^{9c}$ ;
- $R^{9e}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl, Cl, F, Br, I, CN,  $NO_2$ ,  $(CF_2)_rCF_3$ ,  $(CH_2)_rOC_{1-5}$  alkyl, OH, SH,  $(CH_2)_rSC_{1-5}$  alkyl,  $(CH_2)_rNR^{9f}R^{9f}$ , and  $(CH_2)_r$ phenyl, wherein the phenyl on the  $(CH_2)_r$ phenyl is substituted with 0-5 substituents selected from F, Cl, Br, I,  $NO_2$ ,  $C_{1-6}$  alkyl, OH, and  $NR^{9f}R^{9f}$ ;
- $R^{9f}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- $R^{9g}$  is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl,  $(CH_2)_r$ phenyl,  $C(0)R^{9f}$ ,  $C(0)OR^{9h}$ , and  $SO_2R^{9h}$ ;
- $R^{9h}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- $R^{10}$ , is selected from H,  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl, F, Cl, Br, I,  $NO_2$ , CN,  $(CHR')_rOH$ ,

- $\begin{array}{l} (\text{CH}_2)_r \text{OR}^{10d}, \ (\text{CH}_2)_r \text{SR}^{10d}, \ (\text{CH}_2)_r \text{NR}^{10a} \text{R}^{10a'}, \\ (\text{CH}_2)_r \text{C}(0) \text{OH}, \ (\text{CH}_2)_r \text{C}(0) \text{R}^{10b}, \ (\text{CH}_2)_r \text{C}(0) \text{NR}^{10a} \text{R}^{10a'}, \\ (\text{CH}_2)_r \text{NR}^{10a} \text{C}(0) \text{R}^{10a}, \ (\text{CH}_2)_r \text{NR}^{10a} \text{C}(0) \text{H}, \\ (\text{CH}_2)_r \text{C}(0) \text{OR}^{10b}, \ (\text{CH}_2)_r \text{OC}(0) \text{R}^{10b}, \\ (\text{CH}_2)_r \text{OC}(0) \text{NR}^{10a} \text{R}^{10a'}, \ (\text{CH}_2)_r \text{NR}^{10a} \text{C}(0) \text{OR}^{10b}, \\ (\text{CH}_2)_r \text{S}(0)_p \text{R}^{10b}, \ (\text{CH}_2)_r \text{S}(0)_2 \text{NR}^{10a} \text{R}^{10a'}, \\ (\text{CH}_2)_r \text{NR}^{10a} \text{S}(0)_2 \text{R}^{10b}, \ \text{C}_{1-6} \ \text{haloalkyl}, \ \text{a} \ (\text{CH}_2)_r \text{-C}_{3-10} \\ \text{carbocyclic residue substituted with } 0-5 \ \text{R}^{10c}, \ \text{and} \\ \text{a} \ (\text{CH}_2)_r \text{-5-10} \ \text{membered heterocyclic system} \\ \text{containing } 1-4 \ \text{heteroatoms selected from N, O, and} \\ \text{S, substituted with } 0-3 \ \text{R}^{10c}; \\ \end{array}$
- $R^{10a}$  and  $R^{10a'}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r-C_{3-10}$  carbocyclic residue substituted with 0-5  $R^{10e}$ , and a  $(CH_2)_r-5-10$  membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{10e}$ ;
- alternatively, R<sup>10a</sup> and R<sup>10a'</sup>, along with the N to which they are attached, jointo form a 5-6 membered heterocyclic system containing 1-2 heteroatoms selected from NR<sup>10g</sup>, O, and S and optionally fused with a benzene ring or a 6-membered aromatic heterocycle;
- $R^{10b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r$ - $C_{3-6}$  carbocyclic residue substituted with 0-2  $R^{10e}$ , and a  $(CH_2)_r$ -5-6 membered heterocyclic system

containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{10e}$ ;

- R10c, at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl,  $C_{1}$ , Br, I, F,  $(CF_2)_rCF_3$ ,  $NO_2$ , CN,  $(CH_2)_rNR^{10f}R^{10f}$ ,  $(CH_2)_rOH$ ,  $(CH_2)_rOR^{10b}$ ,  $(CH_2)_rSR^{10b}$ ,  $(CH_2)_rC(O)OH$ ,  $(CH_2)_rC(O)R^{10b}$ ,  $(CH_2)_rC(O)NR^{10f}R^{10f}$ ,  $(CH_2)_rNR^{10f}C(O)R^{10a}$ ,  $(CH_2)_rC(O)OR^{10b}$ ,  $(CH_2)_rC(O)OR^{10b}$ ,  $(CH_2)_rC(O)R^{10f}NR^{10f}R^{10f}$ ,  $(CH_2)_rS(O)_pR^{10b}$ ,  $(CH_2)_rNHC(=NR^{10f})NR^{10f}R^{10f}$ ,  $(CH_2)_rS(O)_2NR^{10f}R^{10f}$ ,  $(CH_2)_rNR^{10f}S(O)_2R^{10b}$ , and  $(CH_2)_r$ phenyl substituted with 0-3  $R^{10e}$ ;
- $R^{10d}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  alkenyl,  $C_{3-6}$  alkynyl, and a  $C_{3-10}$  carbocyclic residue substituted with 0-3  $R^{10c}$ ;
- R10e, at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl, Cl, F, Br, I, CN, NO<sub>2</sub>,  $(CF_2)_rCF_3$ ,  $(CH_2)_rOC_{1-5}$  alkyl, OH, SH,  $(CH_2)_rSC_{1-5}$  alkyl,  $(CH_2)_rNR^{10f}R^{10f}$ , and  $(CH_2)_r$ phenyl;
- $R^{10\,f}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- $R^{10g}$  is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl,  $(CH_2)_r$ phenyl,  $C(O)R^{10f}$ ,  $SO_2R^{10h}$ , and  $C(O)OR^{10h}$ ;

- $R^{10h}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl;
- alternatively, R<sup>9</sup> and R<sup>10</sup> join to form =0, a C<sub>3-10</sub> cycloalkyl, a 5-6-membered lactone or lactam, or a 4-6-membered saturated heterocycle containing 1-2 heteroatoms selected from 0, S, and NR<sup>10g</sup> and optionally fused with a benzene ring or a 6-membered aromatic heterocycle;
- with the proviso that when either of R<sup>9</sup> or R<sup>10</sup> is bonded to the carbon to which it is attached through a heteroatom, then the other of R<sup>9</sup> or R<sup>10</sup> is not halogen, cyano, or bonded to the carbon to which it is attached through a heteroatom;
- R11, is selected from H,  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CR'R^{17})_qOH$ ,  $(CH_2)_qSH$ ,  $(CR'R^{17})_qOR^{11d}$ ,  $(CH_2)_qSR^{11d}$ ,  $(CR'R^{17})_qNR^{11a}R^{11a'}$ ,  $(CH_2)_rC(O)OH$ ,  $(CH_2)_rC(O)R^{11b}$ ,  $(CH_2)_rC(O)NR^{11a}R^{11a'}$ ,  $(CH_2)_qNR^{11a}C(O)R^{11a}$ ,  $(CH_2)_qOC(O)NR^{11a}R^{11a'}$ ,  $(CH_2)_qNR^{11a}C(O)OR^{11b}$ ,  $(CH_2)_qNR^{11a}C(O)NHR^{11a}$ ,  $(CH_2)_rC(O)OR^{11b}$ ,  $(CH_2)_qOC(O)R^{11b}$ ,  $(CH_2)_qS(O)_pR^{11b}$ ,  $(CH_2)_qS(O)_2NR^{11a}R^{11a'}$ ,  $(CH_2)_qNR^{11a}S(O)_2R^{11b}$ ,  $C_{1-6}$  haloalkyl, a  $(CH_2)_r-C_{3-10}$  carbocyclic residue substituted with 0-5  $R^{11c}$ , and a  $(R'R^{17})_r-5-10$  membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{11c}$ ;

- Rlla and Rlla', at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r-C_{3-10}$  carbocyclic residue substituted with 0-5  $R^{11e}$ , and a  $(CH_2)_r-5-10$  membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{11e}$ ;
- alternatively, R<sup>11a</sup> and R<sup>11a</sup> along with the N to which they are attached, join to jointe form a 5-6 membered heterocyclic system containing 1-2 heteroatoms selected from NR<sup>11g</sup>, O, and S and optionally fused with a benzene ring or a 6-membered aromatic heterocycle;
- R<sup>11b</sup>, at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r$ - $C_{3-6}$  carbocyclic residue substituted with 0-2 R<sup>11e</sup>, and a  $(CH_2)_r$ -5-6 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3 R<sup>11e</sup>;
- R11c, at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl, Cl, Br, I, F,  $(CF_2)_rCF_3$ ,  $NO_2$ , CN,  $(CH_2)_rNR^{11f}R^{11f}$ ,  $(CH_2)_rOH$ ,  $(CH_2)_rOC_{1-4}$  alkyl,  $(CH_2)_rSC_{1-4}$  alkyl,  $(CH_2)_rC(0)OH$ ,  $(CH_2)_rC(0)R^{11b}$ ,  $(CH_2)_rC(0)NR^{11f}R^{11f}$ ,  $(CH_2)_rNR^{11f}C(0)R^{11a}$ ,  $(CH_2)_rC(0)OC_{1-4}$  alkyl,  $(CH_2)_rOC(0)R^{11b}$ ,  $(CH_2)_rC(0)NR^{11f}R^{11f}$ ,  $(CH_2)_rOC(0)R^{11b}$ ,  $(CH_2)_rC(0)NR^{11f}R^{11f}$ ,  $(CH_2)_rNHC(0)R^{11f}NR^{11f}R^{11f}$ ,  $(CH_2)_rNHC(0)R^{11f}NR^{11f}R^{11f}$ ,  $(CH_2)_rNHC(0)R^{11f}NR^{11f}R^{11f}$ ,  $(CH_2)_rS(0)_pR^{11b}$ ,

- $(CH_2)_rS(O)_2NR^{11f}R^{11f}$ ,  $(CH_2)_rNR^{11f}S(O)_2R^{11b}$ , and  $(CH_2)_r$ phenyl substituted with 0-3  $R^{11e}$ ;
- $R^{11d}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6} \ \, \text{alkenyl}, \ \, C_{3-6} \ \, \text{alkynyl}, \ \, \text{and a} \ \, C_{3-10} \ \, \text{carbocyclic}$  residue substituted with 0-3  $R^{11c}$ ;
- R<sup>11e</sup>, at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $C_{3-6}$  cycloalkyl, Cl, F, Br, I, CN, NO<sub>2</sub>,  $(CF_2)_rCF_3$ ,  $(CH_2)_rOC_{1-5}$  alkyl, OH, SH,  $(CH_2)_rSC_{1-5}$  alkyl,  $(CH_2)_rNR^{11f}R^{11f}$ , and  $(CH_2)_r$ phenyl, wherein the phenyl on the  $(CH_2)_r$ phenyl is substituted with 0-5 substituents selected from F, Cl, Br, I, NO<sub>2</sub>,  $C_{1-6}$ alkyl, OH, and  $NR^{9f}R^{9f}$ ;
- $R^{11f}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- $R^{11g}$  is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl,  $(CH_2)_r$ phenyl,  $C(0)R^{11f}$ ,  $C(0)OR^{11h}$ , and  $SO_2R^{11h}$ ;
- $R^{11h}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- R<sup>12</sup>, is selected from H, C<sub>1-6</sub> alkyl, C<sub>2-8</sub> alkenyl, C<sub>2-8</sub> alkynyl, (CHR')<sub>q</sub>OH, (CH<sub>2</sub>)<sub>q</sub>SH, (CHR')<sub>q</sub>OR<sup>12d</sup>, (CH<sub>2</sub>)<sub>q</sub>SR<sup>12d</sup>, (CHR')<sub>q</sub>NR<sup>12a</sup>R<sup>12a'</sup>, (CH<sub>2</sub>)<sub>r</sub>C(O)OH, (CH<sub>2</sub>)<sub>r</sub>C(O)R<sup>12b</sup>, (CH<sub>2</sub>)<sub>r</sub>C(O)NR<sup>12a</sup>R<sup>12a'</sup>,

 $\begin{array}{l} (\text{CH}_2)_q \text{NR}^{12a} \text{C}(0) \, \text{R}^{12a}, & (\text{CH}_2)_r \text{OC}(0) \, \text{NR}^{12a} \text{R}^{12a'}, \\ (\text{CH}_2)_r \text{NR}^{12a} \text{C}(0) \, \text{OR}^{12b}, & (\text{CH}_2)_q \text{NR}^{12a} \text{C}(0) \, \text{NHR}^{12a}, \\ (\text{CH}_2)_r \text{C}(0) \, \text{OR}^{12b}, & (\text{CH}_2)_q \text{OC}(0) \, \text{R}^{12b}, & (\text{CH}_2)_q \text{S}(0)_p \text{R}^{12b}, \\ (\text{CH}_2)_q \text{S}(0)_2 \text{NR}^{12a} \text{R}^{12a'}, & (\text{CH}_2)_q \text{NR}^{12a} \text{S}(0)_2 \text{R}^{12b}, & \text{C}_{1-6} \\ \text{haloalkyl, a} & (\text{CH}_2)_r - \text{C}_{3-10} & \text{carbocyclic residue} \\ \text{substituted with 0-5} \, \text{R}^{12c}, & \text{and a} & (\text{R}'\text{R}^{17})_r - \text{5-10} \\ \text{membered heterocyclic system containing 1-4} \\ \text{heteroatoms selected from N, 0, and S, substituted} \\ \text{with 0-3} \, \text{R}^{12c}; \end{array}$ 

- $R^{12a}$  and  $R^{12a'}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r-C_{3-10}$  carbocyclic residue substituted with 0-5  $R^{12e}$ , and a  $(CH_2)_r-5-10$  membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{12e}$ ;
- alternatively, R<sup>12a</sup> and R<sup>12a</sup>, along with the N to which they are attached, jointo form a 5-6 membered heterocyclic system containing 1-2 heteroatoms selected from NR<sup>12g</sup>, O, and S and optionally fused with a benzene ring or a 6-membered aromatic heterocycle;
- $R^{12b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r$ - $C_{3-6}$  carbocyclic residue substituted with 0-2  $R^{12e}$ , and a  $(CH_2)_r$ -5-6 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{12e}$ ;

- R12c, at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl, Cl, Br, I, F,  $(CF_2)_rCF_3$ ,  $NO_2$ , CN,  $(CH_2)_rNR^{12f}R^{12f}$ ,  $(CH_2)_rOH$ ,  $(CH_2)_rOC_{1-4}$  alkyl,  $(CH_2)_rSC_{1-4}$  alkyl,  $(CH_2)_rC(0)OH$ ,  $(CH_2)_rC(0)R^{12b}$ ,  $(CH_2)_rC(0)NR^{12f}R^{12f}$ ,  $(CH_2)_rNR^{12f}C(0)R^{12a}$ ,  $(CH_2)_rC(0)OC_{1-4}$  alkyl,  $(CH_2)_rOC(0)R^{12b}$ ,  $(CH_2)_rC(0)OC_{1-4}$  alkyl,  $(CH_2)_rOC(0)R^{12b}$ ,  $(CH_2)_rC(0)NR^{12f}R^{12f}$ ,  $(CH_2)_rNHC(=NR^{12f})NR^{12f}R^{12f}$ ,  $(CH_2)_rS(0)_pR^{12b}$ ,  $(CH_2)_rS(0)_2NR^{12f}R^{12f}$ ,  $(CH_2)_rNR^{12f}S(0)_2R^{12b}$ , and  $(CH_2)_r$ phenyl substituted with 0-3  $R^{12e}$ ;
- $R^{12d}$ , at each occurrence, is selected from methyl,  $CF_3$ ,  $C_{2-6}$  alkyl substituted with 0-3  $R^{12e}$ ,  $C_{3-6}$  alkenyl, and a  $C_{3-10}$  carbocyclic residue substituted with 0-3  $R^{12c}$ ;
- R<sup>12f</sup>, at each occurrence, is selected from H, C<sub>1-6</sub> alkyl, and C<sub>3-6</sub> cycloalkyl;
- $R^{12g}$  is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl,  $(CH_2)_r$ phenyl,  $C(O)R^{12f}$ ,  $C(O)OR^{12h}$ , and  $SO_2R^{12h}$ ;

- $R^{12h}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- alternatively, R<sup>11</sup> and R<sup>12</sup> join to form a C<sub>3-10</sub> cycloalkyl, a 5-6-membered lactone or lactam, or a 4-6-membered saturated heterocycle containing 1-2 heteroatoms selected from O, S, and NR<sup>11g</sup> and optionally fused with a benzene ring or a 6-membered aromatic heterocycle;
- R<sup>13</sup>, at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $C_{3-6}$  cycloalkyl,  $(CF_2)_w CF_3, \ (CH_2)_q NR^{13a}R^{13a'}, \ (CHR')_q OH, \ (CH_2)_q OR^{13b}, \\ (CH_2)_q SH, \ (CH_2)_q SR^{13b}, \ (CH_2)_w C(0) OH, \ (CH_2)_w C(0) R^{13b}, \\ (CH_2)_w C(0) NR^{13a}R^{13a'}, \ (CH_2)_q NR^{13d}C(0)R^{13a}, \\ (CH_2)_w C(0) OR^{13b}, \ (CH_2)_q OC(0)R^{13b}, \ (CH_2)_w S(0)_p R^{13b}, \\ (CH_2)_w S(0)_2 NR^{13a}R^{13a'}, \ (CH_2)_q NR^{13d}S(0)_2 R^{13b}, \ and \\ (CH_2)_w -phenyl \ substituted \ with \ 0-3 \ R^{13c};$
- $R^{13a}$  and  $R^{13a}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl substituted with 0-3  $R^{13c}$ ;
- $R^{13b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl substituted with 0-3  $R^{13c}$ .
- $R^{13c}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, Cl, F, Br, I, CN,  $NO_2$ ,  $(CF_2)_rCF_3$ ,

- $(CH_2)_rOC_{1-5}$  alkyl,  $(CH_2)_rOH$ ,  $(CH_2)_rSC_{1-5}$  alkyl, and  $(CH_2)_rNR^{13}dR^{13}d$ ;
- $R^{13d}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- R14, at each occurrence, is selected from H, C1-6 alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$ cycloalkyl, Cl, Br, I, F, NO2, CN,  $(CHR')_rNR^{14a}R^{14a'}$ ,  $(CHR')_rOH$ ,  $(CHR')_rO(CHR')_rR^{14d}$ ,  $(CHR')_rSH$ ,  $(CHR')_rC(O)H$ ,  $(CHR')_rS(CHR')_rR^{14d}$ ,  $(CHR')_rC(O)OH$ ,  $(CHR')_rC(O)(CHR')_rR^{14b}$ ,  $(CHR')_rC(O)NR^{14a}R^{14a'}$ ,  $(CHR')_rNR^{14f}C(O)(CHR')_rR^{14b}$ ,  $(CHR')_rOC(O)NR^{14a}R^{14a'}, (CHR')_rNR^{14f}C(O)O(CHR')_rR^{14b},$  $(CHR')_rC(O)O(CHR')_rR^{14d}$ ,  $(CHR')_rOC(O)(CHR')_rR^{14b}$ ,  $(CHR')_rC (=NR^{14f})NR^{14a}R^{14a'}$ ,  $(CHR')_rNHC (=NR^{14f})NR^{14f}R^{14f}, (CHR')_rS(O)_p(CHR')_rR^{14b},$  $(CHR')_rS(O)_2NR^{14a}R^{14a'}, (CHR')_rNR^{14f}S(O)_2(CHR')_rR^{14b},$  $C_{1-6}$  haloalkyl,  $C_{2-8}$  alkenyl substituted with 0-3 R',  $C_{2-8}$  alkynyl substituted with 0-3 R', (CHR') phenyl substituted with 0-3  $\rm R^{14e}$ , and a (CH<sub>2</sub>)<sub>r</sub>-5-10 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-2 R15e, or two R14 substituents on adjacent atoms on ring A form to join a 5-6 membered heterocyclic system containing 1-3 heteroatoms selected from N, O, and S substituted with 0-2 R<sup>15e</sup>;

- $R^{14a}$  and  $R^{14a'}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r-C_{3-10}$  carbocyclic residue substituted with 0-5  $R^{14e}$ , and a  $(CH_2)_r-5-10$  membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-2  $R^{14e}$ ;
- $R^{14b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r$ - $C_{3-6}$  carbocyclic residue substituted with 0-3  $R^{14e}$ , and  $(CH_2)_r$ -5-6 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-2  $R^{14e}$ ;
- R14d, at each occurrence, is selected from  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, methyl,  $CF_3$ ,  $C_{2-6}$  alkyl substituted with 0-3  $R^{14e}$ , a  $(CH_2)_r$ - $C_{3-10}$  carbocyclic residue substituted with 0-3  $R^{14e}$ , and a  $(CH_2)_r$ 5-6 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{14e}$ ;
- R<sup>14e</sup>, at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl, Cl, F, Br, I, CN, NO<sub>2</sub>,  $(CF_2)_rCF_3$ ,  $(CH_2)_rOC_{1-5}$  alkyl, OH, SH,  $(CH_2)_rSC_{1-5}$  alkyl,  $(CH_2)_rNR^{14f}R^{14f}$ , and  $(CH_2)_rphenyl$ ;
- $R^{14f}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl;

- $R^{15}$ , at each occurrence, is selected from  $C_{1-8}$  alkyl, (CH<sub>2</sub>)<sub>r</sub>C<sub>3-6</sub> cycloalkyl, Cl, Br, I, F, NO<sub>2</sub>, CN,  $(CR'R^{17})_rNR^{15a}R^{15a'}, (CR'R^{17})_rOH,$  $(CR'R^{17})_rO(CHR')_rR^{15d}$ ,  $(CR'R^{17})_rSH$ ,  $(CR'R^{17})_rC(O)H$ ,  $(CR'R^{17})_rS(CHR')_rR^{15d}$ ,  $(CR'R^{17})_rC(O)OH$ ,  $(CR'R^{17})_rC(0)(CHR')_rR^{15b}, (CR'R^{17})_rC(0)NR^{15a}R^{15a'},$  $(CR'R^{17})_rNR^{15f}C(0)(CHR')_rR^{15b},$  $(CR'R^{17})_rOC(O)NR^{15a}R^{15a'}$ ,  $(CR'R^{17})_rNR^{15f}C(0)O(CHR')_rR^{15b}$ ,  $(CR'R^{17})_rNR^{15f}C(0)NR^{15f}R^{15f}$ ,  $(CR'R^{17})_rC(0)O(CHR')_rR^{15d}$ ,  $(CR'R^{17})_rOC(O)(CHR')_rR^{15b}$ ,  $(CR'R^{17})_rC(=NR^{15f})NR^{15a}R^{15a'}$ ,  $(CR'R^{17})_rNHC (=NR^{15f})NR^{15f}R^{15f}$  $(CR'R^{17})_rS(0)_p(CHR')_rR^{15b}, (CR'R^{17})_rS(0)_2NR^{15a}R^{15a'},$  $(CR'R^{17})_rNR^{15f}S(0)_2(CHR')_rR^{15b}, C_{1-6} haloalkyl, C_{2-8}$ alkenyl substituted with 0-3 R', C2-8 alkynyl substituted with 0-3 R', (CR'R<sup>17</sup>)<sub>r</sub>phenyl substituted with 0-3  $R^{15e}$ , and a  $(CH_2)_r$ -5-10 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with  $0-2 R^{15e}$ ;
- $R^{15a}$  and  $R^{15a'}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r$ - $C_{3-10}$  carbocyclic residue substituted with 0-5  $R^{15e}$ , and a  $(CH_2)_r$ -5-10 membered heterocyclic system

- containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-2  $R^{15e}$ ;
- alternatively, R<sup>15a</sup> and R<sup>15a'</sup>, along with the N to which they are attached, join to jointe form a 5-6 membered heterocyclic system containing 1-2 heteroatoms selected from NR<sup>15h</sup>, O, and S and optionally fused with a benzene ring or a 6-membered aromatic heterocycle;
- $R^{15b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r-C_{3-6}$  carbocyclic residue substituted with 0-3  $R^{15e}$ , and  $(CH_2)_r-5-6$  membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-2  $R^{15e}$ ;
- $R^{15d}$ , at each occurrence, is selected from  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, methyl,  $CF_3$ ,  $C_{2-6}$  alkyl substituted with 0-3  $R^{15e}$ , a  $(CH_2)_r$ - $C_{3-10}$  carbocyclic residue substituted with 0-3  $R^{15e}$ , and a  $(CH_2)_r$ 5-6 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{15e}$ ;
- $R^{15e}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl, 2-cyanoethyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl, Cl, F, Br, I, CN,  $NO_2$ ,  $(CF_2)_rCF_3$ ,  $(CH_2)_rOC_{1-5}$  alkyl, OH, SH,  $(CH_2)_rSC_{1-5}$  alkyl,  $(CH_2)_rNR^{15f}R^{15f}$ ,  $(CH_2)_rphenyl$ , and a heterocycle

substituted with 0-1 R<sup>15g</sup>, wherein the heterocycle is selected from imidazole, thiazole, oxazole, pyrazole, 1,2,4-triazole, 1,2,3-triazole, isoxazole, and tetrazole,;

- $R^{15f}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl;
- $R^{15g}$  is selected from methyl, ethyl, acetyl, and  $CF_3$ ;
- $R^{15h}$  is selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl,  $(CH_2)_r$ phenyl,  $C(O)R^{15f}$ ,  $C(O)OR^{15i}$ , and  $SO_2R^{15i}$ ;
- $R^{15i}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl;
- R16, at each occurrence, is selected from  $C_{1-8}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl,  $C_{1}$ , Br, I, F, NO<sub>2</sub>, CN,  $(CHR')_rNR^{16a}R^{16a'}$ ,  $(CHR')_rOH$ ,  $(CHR')_rO(CHR')_rR^{16d}$ ,  $(CHR')_rSH$ ,  $(CHR')_rC(O)H$ ,  $(CHR')_rS(CHR')_rR^{16d}$ ,  $(CHR')_rC(O)OH$ ,  $(CHR')_rC(O)(CHR')_rR^{16b}$ ,  $(CHR')_rC(O)NR^{16a}R^{16a'}$ ,  $(CHR')_rNR^{16f}C(O)(CHR')_rR^{16b}$ ,  $(CHR')_rC(O)O(CHR')_rR^{16b}$ ,  $(CHR')_rC(O)O(CHR')_rR^{16d}$ ,  $(CHR')_rC(O)O(CHR')_rR^{16d}$ ,  $(CHR')_rC(O)O(CHR')_rR^{16b}$ ,  $(CHR')_rC(O)O(CHR^{16f})NR^{16a}R^{16a'}$ ,  $(CHR')_rS(O)_p(CHR')_rR^{16b}$ ,  $(CHR')_rS(O)_2NR^{16a}R^{16a'}$ ,  $(CHR')_rNR^{16f}S(O)_2(CHR')_rR^{16b}$ ,  $(CHR')_rS(O)_2NR^{16a}R^{16a'}$ ,  $(CHR')_rNR^{16f}S(O)_2(CHR')_rR^{16b}$ ,  $(C_{1-6}$  haloalkyl,  $C_{2-8}$  alkenyl substituted with 0-3

- R',  $C_{2-8}$  alkynyl substituted with 0-3 R', and  $(CHR')_r$ phenyl substituted with 0-3 R<sup>16e</sup>;
- $R^{16a}$  and  $R^{16a'}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_r-C_{3-10}$  carbocyclic residue substituted with 0-5  $R^{16e}$ , and a  $(CH_2)_r-5-10$  membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-2  $R^{16e}$ ;
- $R^{16b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, a  $(CH_2)_rC_{3-6}$  carbocyclic residue substituted with 0-3  $R^{16e}$ , and a  $(CH_2)_r$ -5-6 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-2  $R^{16e}$ ;
- $R^{16d}$ , at each occurrence, is selected from  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl, methyl,  $CF_3$ ,  $C_{2-6}$  alkyl substituted with 0-3  $R^{16e}$ , a  $(CH_2)_r$ - $C_{3-10}$  carbocyclic residue substituted with 0-3  $R^{16e}$ , and a  $(CH_2)_r$ -5-6 membered heterocyclic system containing 1-4 heteroatoms selected from N, O, and S, substituted with 0-3  $R^{16e}$ ;
- $R^{16e}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl, Cl, F, Br, I, CN,  $NO_2$ ,  $(CF_2)_rCF_3$ ,  $(CH_2)_rOC_{1-5}$  alkyl, OH, SH,  $(CH_2)_rSC_{1-5}$  alkyl,  $(CH_2)_rNR^{16f}R^{16f}$ , and  $(CH_2)_rphenyl$ ;

- $R^{16f}$ , at each occurrence, is selected from H,  $C_{1-5}$  alkyl, and  $C_{3-6}$  cycloalkyl, and phenyl;
- R<sup>17</sup>, at each occurrence, is independently selected from H and methyl;
- R', at each occurrence, is selected from H,  $C_{1-6}$  alkyl,  $C_{3-8}$  alkenyl,  $C_{3-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl, and  $(CH_2)_r$ phenyl substituted with  $R^{15e}$ ;
- g is selected from 0, 1, 2, 3, and 4;
- v is selected from 0, 1, and 2;
- t is selected from 1 and 2;
- w is selected from 0 and 1;
- r is selected from 0, 1, 2, 3, 4, and 5;
- q is selected from 1, 2, 3, 4, and 5; and
- p is selected from 0, 1, and 2.
  - 2. (ORIGINAL) The compound of claim 1, wherein:
- Z is selected from O, S, N(CN), and N(CONH<sub>2</sub>);
- $R^2$  is selected from H and  $C_{1-4}$  alkyl;

- R<sup>6</sup>, at each occurrence, is selected from  $C_{1-4}$  alkyl,  $C_{2-8}$  alkenyl,  $C_{2-8}$  alkynyl,  $(CH_2)_rC_{3-6}$  cycloalkyl,  $(CF_2)_rCF_3$ , CN,  $(CH_2)_rOH$ ,  $(CH_2)_rOR^{6b}$ ,  $(CH_2)_rC(O)R^{6a}R^{6a'}$ ,  $(CH_2)_rNR^{6d}C(O)R^{6a}$ , and  $(CH_2)_rDhenyl$  substituted with 0-3  $R^{6c}$ ;
- $R^{6a}$  and  $R^{6a'}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl substituted with 0-3  $R^{6c}$ ;
- $R^{6b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl substituted with 0-3  $R^{6c}$ ;
- $R^{6c}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, Cl, F, Br, I, CN,  $NO_2$ ,  $(CF_2)_rCF_3$ ,  $(CH_2)_rOC_{1-5}$  alkyl,  $(CH_2)_rOH$ ,  $(CH_2)_rSC_{1-5}$  alkyl, and  $(CH_2)_rNR^{6d}R^{6d}$ ;
- $R^{6d}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- R<sup>13</sup>, at each occurrence, is selected from  $C_{1-4}$  alkyl,  $C_{3-6}$  cycloalkyl,  $(CH_2)NR^{13a}R^{13a'}$ , (CHR')OH,  $(CH_2)OR^{13b}$ ,  $(CH_2)_wC(O)R^{13b}$ ,  $(CH_2)_wC(O)NR^{13a}R^{13a'}$ ,  $(CH_2)NR^{13d}C(O)R^{13a}$ ,  $(CH_2)_wS(O)_2NR^{13a}R^{13a'}$ ,  $(CH_2)NR^{13d}S(O)_2R^{13b}$ , and  $(CH_2)_w$ -phenyl substituted with 0-3  $R^{13c}$ ;

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- $R^{13a}$  and  $R^{13a}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl substituted with 0-3  $R^{13c}$ ;
- $R^{13b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and phenyl substituted with 0-3  $R^{13c}$ ;
- $R^{13c}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, Cl, F, Br, I, CN,  $NO_2$ ,  $(CF_2)_rCF_3$ ,  $(CH_2)_rOC_{1-5}$  alkyl,  $(CH_2)_rOH$ , and  $(CH_2)_rNR^{13d}R^{13d}$ ;
- $R^{13d}$ , at each occurrence, is selected from H,  $C_{1-6}$  alkyl, and  $C_{3-6}$  cycloalkyl;
- v is selected from 0, 1 and 2;
- q is selected from 1, 2, and 3; and
- r is selected from 0, 1, 2, and 3...
  - 3. (ORIGINAL) The compound of claim 2, wherein:
- E is  $-(C=0)-(CR^9R^{10})_{v}-(CR^{11}R^{12})-$ ,  $-(SO_2)-(CR^9R^{10})_{v}-(CR^{11}R^{12})-$ ,

R<sup>3</sup> is selected from (CH<sub>2</sub>)<sub>2</sub>N(CH<sub>3</sub>)<sub>2</sub>, a (CR<sup>3</sup>'H)<sub>r</sub>carbocyclic residue substituted with 0-5 R<sup>15</sup>,
wherein the carbocyclic residue is selected from
phenyl, C<sub>3-6</sub> cycloalkyl, naphthyl, and adamantyl;
and a (CR<sup>3</sup>'H)<sub>r</sub>-heterocyclic system substituted
with 0-3 R<sup>15</sup>, wherein the heterocyclic system is
selected from pyridinyl, thiophenyl, furanyl,
indazolyl, benzothiazolyl, benzimidazolyl,
benzothiophenyl, benzofuranyl, benzoxazolyl,
benzisoxazolyl, quinolinyl, isoquinolinyl,
imidazolyl, indolyl, indolinyl, isoindolyl,
isothiadiazolyl, isoxazolyl, piperidinyl,
pyrrazolyl, 1,2,4-triazolyl, 1,2,3-triazolyl,
tetrazolyl, thiadiazolyl, thiazolyl, oxazolyl,
pyrazinyl, and pyrimidinyl; and

R<sup>5</sup> is selected from (CR<sup>5</sup>'H)<sub>t</sub>-phenyl substituted with 0-5 R<sup>16</sup>; and a (CR<sup>5</sup>'H)<sub>t</sub>-heterocyclic system substituted with 0-3 R<sup>16</sup>, wherein the heterocyclic system is selected from pyridinyl, thiophenyl, furanyl, indazolyl, benzothiazolyl, benzimidazolyl, benzothiophenyl, benzofuranyl, benzoxazolyl, benzisoxazolyl, quinolinyl, isoquinolinyl, imidazolyl, indolyl, indolinyl,

isoindolyl, isothiadiazolyl, isoxazolyl, piperidinyl, pyrrazolyl, 1,2,4-triazolyl, 1,2,3-triazolyl, tetrazolyl, thiadiazolyl, thiazolyl, oxazolyl, pyrazinyl, and pyrimidinyl.

### 4. (CANCELED)

- # \$. (PREVIOUSLY PRESENTED) The compound of claim 3, wherein
- R<sup>16</sup>, at each occurrence, is selected from C<sub>1-8</sub> alkyl,  $(CH_2)_r C_{3-6} \text{ cycloalkyl}, CF_3, Cl, Br, I, F, \\ (CH_2)_r NR^{16a}R^{16a'}, NO_2, CN, OH, (CH_2)_r OR^{16d}, \\ (CH_2)_r C(O)R^{16b}, (CH_2)_r C(O)NR^{16a}R^{16a'}, \\ (CH_2)_r NR^{16f}C(O)R^{16b}, (CH_2)_r S(O)_p R^{16b}, \\ (CH_2)_r S(O)_2 NR^{16a}R^{16a'}, (CH_2)_r NR^{16f}S(O)_2 R^{16b}, \text{ and} \\ (CH_2)_r Dhenyl substituted with 0-3 R^{16e};$
- $R^{16a}$  and  $R^{16a'}$ , at each occurrence, are selected from H,  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and  $(CH_2)_r$ phenyl substituted with 0-3  $R^{16e}$ ;
- $R^{16b}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl,  $C_{3-6}$  cycloalkyl, and  $(CH_2)_r$ phenyl substituted with 0-3  $R^{16e}$ ;
- $R^{16d}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl and phenyl;

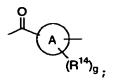
 $R^{16e}$ , at each occurrence, is selected from  $C_{1-6}$  alkyl, Cl, F, Br, I, CN,  $NO_2$ ,  $(CF_2)_rCF_3$ , OH, and  $(CH_2)_rOC_{1-5}$  alkyl; and

 $R^{16f}$ , at each occurrence, is selected from H, and  $C_{1-5}$  alkyl.

### 6. (CANCELED)

f. (ORIGINAL) The compound of claim 3, wherein:

E is  $-(C=0)-(CR^9R^{10})_{v}-(CR^{11}R^{12})-$ , or



 $R^5$  is CH2phenyl substituted with 0-3  $R^{16}$ ; and

r is selected from 0, 1, and 2.

### 8. (CANCELED)

6 8. (ORIGINAL) The compound of claim 1, wherein:

K is selected from CH<sub>2</sub> and CHR<sup>5</sup>;

L is selected from CH<sub>2</sub> and CHR<sup>5</sup>; and

- R<sup>3</sup> is a (CH<sub>2</sub>)<sub>r</sub>-C<sub>3-10</sub> carbocyclic residue substituted with 0-3 R<sup>15</sup>, wherein the carbocyclic residue is selected from cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl, phenyl, naphthyl and adamantyl, and a (CR<sup>3</sup>'H)<sub>r</sub>-heterocyclic system substituted with 0-3 R<sup>15</sup>, wherein the heterocyclic system is selected from pyridinyl, thiophenyl, furanyl, indazolyl, benzothiazolyl, benzimidazolyl, benzothiophenyl, benzofuranyl, benzoxazolyl, benzisoxazolyl, quinolinyl, isoquinolinyl, imidazolyl, indolyl, indolyl, indolyl, isoindolyl, isothiadiazolyl, isoxazolyl, piperidinyl, pyrrazolyl, 1,2,4-triazolyl, 1,2,3-triazolyl, tetrazolyl, thiadiazolyl, thiazolyl, oxazolyl, pyrazinyl, and pyrimidinyl.
  - 7 10. (PREVIOUSLY PRESENTED) The compound of claim 3, wherein:

K and L are independently selected from  $CH_2$  and  $CHR^5$ ;

Z is O, S, NCN, or NCONH<sub>2</sub>;

 $\mathbb{R}^1$  is H;

 $R^2$  is H:

 $R^3$  is selected from a  $(CH_2)_rN(CH_3)_2$ , a  $(CH_2)_r-C_{3-10}$  carbocyclic residue substituted with 0-3  $R^{15}$ , wherein the carbocyclic residue is selected from cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl,

phenyl, naphthyl and adamantyl, and a (CR3'H)<sub>r</sub>-heterocyclic system substituted with 0-3 R<sup>15</sup>, wherein the heterocyclic system is selected from pyridinyl, thiophenyl, furanyl, indazolyl, benzothiazolyl, benzimidazolyl, benzothiophenyl, benzofuranyl, benzoxazolyl, benzisoxazolyl, quinolinyl, isoquinolinyl, imidazolyl, indolyl, indolinyl, isoindolyl, isothiadiazolyl, isoxazolyl, piperidinyl, pyrrazolyl, 1,2,4-triazolyl, 1,2,3-triazolyl, tetrazolyl, thiadiazolyl, and pyrimidinyl; and

R<sup>5</sup> is selected from a CH<sub>2</sub>-phenyl substituted with 0-5
R<sup>16</sup> and a CH<sub>2</sub>-heterocyclic system substituted with
0-3 R<sup>16</sup>, wherein the heterocyclic system is
selected from pyridinyl, thiophenyl, furanyl,
indazolyl, benzothiazolyl, benzimidazolyl,
benzothiophenyl, benzofuranyl, benzoxazolyl,
benzisoxazolyl, quinolinyl, isoquinolinyl,
imidazolyl, indolyl, indolinyl, isoindolyl,
isothiadiazolyl, isoxazolyl, piperidinyl,
pyrrazolyl, 1,2,4-triazolyl, 1,2,3-triazolyl,
tetrazolyl, thiadiazolyl, thiazolyl, oxazolyl,
pyrazinyl, and pyrimidinyl.

#### 11. (CANCELED)

### 12. (CANCELED)

(ORIGINAL) A pharmaceutical composition, comprising a pharmaceutically acceptable carrier and a

therapeutically effective amount of a compound according to Claim 1.

#### 14. (CANCELLED)

- <sup>c</sup> 15. (PREVIOUSLY PRESENTED) A method for treating asthma, comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to Claim 1.
- 10 16. (ORIGINAL) A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound according to Claim 1, or a pharmaceutically acceptable salt thereof.
  - 17. (CANCELLED)
  - 18. (CANCELLED)
- inflammation in an inflammatory disorders comprising administering to a patient in need thereof a therapeutically effective amount of a compound according to Claim 42, or a pharmaceutically acceptable salt thereof A method according to Claim 18, wherein the disorder is selected from asthma, allergic rhinitis, atopic dermatitis, inflammatory bowel diseases, idiopathic pulmonary fibrosis, bullous pemphigoid, helminthic parasitic infections, allergic colitis, eczema, conjunctivitis, transplantation, familial eosinophilia, eosinophilic cellulitis,

eosinophilic pneumonias, eosinophilic fasciitis, <u>and</u> eosinophilic gastroenteritis, <u>drug induced</u> eosinophilia, HIV infection, cystic fibrosis, Churg-Strauss syndrome, lymphoma, Hodgkin's disease, and colonic carcinoma.

- 20. (ORIGINAL) The method according to Claim 19, wherein the disorder is selected from asthma, allergic rhinitis, atopic dermatitis, and inflammatory bowel diseases.
- 21. (ORIGINAL) The method according to Claim 20, wherein the disorder is asthma.

Respectfully submitted,

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